## AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listings, of claims in the application:

- 1. (original) Non-human transgenic animal, being transgenic for having altered melusin expression.
- 2. (original) Non-human transgenic animal according to claim 1, characterized in that said altered melusin expression is performed by stable or transient modification of melusin expression at transcriptional, translational or post- translational level.
- 3. (currently amended) Non-human transgenic animal according to claim 1 or 2, characterized in that said altered melusin expression is an inactivation of melusin gene.
- 4. (original) Non-human transgenic animal according to claim 3, characterized in that said gene inactivation is performed by genetic approaches.
- 5. (original) Non-human transgenic animal according to claim 4, characterized in that said genetic approaches are selected from the group consisting of homologous recombination, antisense RNA or DNA and RNA or DNA interference approach.
- 6. (currently amended) Non-human transgenic animal according to any of the preceding claims claim 1, characterized in that said animal is a melusin-null transgenic animal.
- 7. (currently amended) Non-human transgenic animal according to any of the claims 1 to 6 claim 1, characterized in that said animal is subjected to hypertensive condition.
- 8. (original) Non-human transgenic animal according to claim 7, characterized in that said hypertensive condition is determined by surgical operation.
- 9. (original) Non-human transgenic animal according to claim 8, characterized in that said surgical operation consists in surgical constriction of the transverse aorta.

## BRANCACCIO et al U.S. National Phase of PCT/IT2002/000807

- 10. (original) Non-human transgenic animal according to claim 7, characterized in that said hypertensive condition is determined by pharmacological treatment, preferably with hypertensive drugs.
- 11. (original) Non-human transgenic animal according to claim 7, characterized in that said hypertensive condition is determined by high sodium diet.
- 12. (currently amended) Non-human transgenic animal according any of the preceding claims to claim 1, wherein said animal develop at least impaired heart hypertrophy.
- 13. (currently amended) Non-human transgenic animal according any of the preceding claims to claim 1, wherein said animal develop at least heart dilation.
- 14. (currently amended) Non-human transgenic animal according any of the preceding claims to claim 1, wherein said animal develop at least heart failure.
- 15. (currently amended) Non-human transgenic animal according any of the preceding claims to claim 1, wherein said animal is a mammalian.
- 16. (original) Non-human transgenic animal according to claim 15, wherein the mammalian belongs to the murine genus (mus musculus).
- 17. (original) Non-human transgenic animal according to claim 16, wherein said mouse belongs to the 129SV, C57B1 or 129SVxC57BI strain.
- 18. (currently amended) Use of non-human transgenic animal according to any of the preceding claims claim 1 for the selection of compounds pharmacologically active in the prevention and/or treatment of heart failure.
- 19. (currently amended) Use of non-human transgenic animal according to any of claims 1 to 17 claim 1 for the study of heart pathologies, wherein said heart pathologies are selected from the group consisting of: heart failure, congestive heart failure, dilated cardiomyopathy, hypertensive cardiomyopathy, hypertrophic cardiomyopathy, heart infarct.

## BRANCACCIO et al U.S. National Phase of PCT/IT2002/000807

- 20. (currently amended) Cells derivable from the non-human transgenic animal according to any of claims 1 to 17 claim 1 and having altered melusin expression.
- 21. (original) Cells according to claim 20, characterized in that said cells carry a mutation inactivating melusin gene.
- 22. (currently amended) Cells according to claim 20 or 21, characterized in that said cells are lacking melusin expression.
- 23. (currently amended) Use of cells according to any claim 20 to 22 claim 20 for the screening of compounds pharmacologically active for the prevention and/or treatment of heart failure.
- 24. (original) Method for the preparation of a non-human transgenic animal according to claim 1 comprising essentially the steps of:
- i) preparing a non-human transgenic parent animal carrying an inactivated melusin allele;
  - ii) breeding the parent transgenic animal with a non transgenic animal;
  - iii) selecting transgenic animals heterogyzote for the melusin gene mutation.
- 25. (original) Method according to claim 24, further comprising the step of iv) breeding the heterozygote transgenic animals to select homozygote transgenic animals for the melusin gene mutation.
- 26. (original) Non-human animals in which melusin function has been inhibited by the use of natural or synthetic compounds.
- 27. (original) Use of the animal according to claim 26 to study the impaired cardiac hypertrophy.
  - 28. (original) Use of the animal according to claim 26 to study cardiac dilation.
  - 29. (original) Use of the animal according to claim 26 to study the heart failure.
- 30. (original) Method for screening compounds able to interact with melusin binding proteins, said compounds being pharmacologically active in the prevention

## BRANCACCIO et al U.S. National Phase of PCT/IT2002/000807

and/or treatment of heart failure, wherein said method comprises using melusin, fragments and/or derivatives thereof.

- 31. (original) Method for screening compounds able to interact with melusin, said compounds being melusin agonists and pharmacologically active in the prevention and/or treatment of heart failure, wherein said method comprises using melusin, fragments and/or derivatives thereof.
- 32. (original) Use of melusin, fragments and/or derivatives thereof for the manufacture of a medicament for the prevention and/or treatment of heart failure.
- 33. (original) Use of melusin, fragments and/or derivatives thereof for the screening of compounds pharmacologically active for the prevention and/or treatment of heart failure.
- 34. (original) Use according to claim 33, characterized in that said pharmacologically active compound is a melusin agonist.
- 35. (original) Use according to claim 33, characterized in that said pharmacologically active compound is able to interact with melusin-binding proteins.
- 36. (original) Use of a DNA vector for the manufacture of a medicament for use in the prevention and/or treatment of heart failure, said vector comprising a transgene coding for the melusin protein or fragments thereof and expressing said transgene in the myocardium.
- 37. (original) Use according to claim 36, characterized in that said transgene comprises melusin cDNA or fragments thereof.
- 38. (currently amended) Use according to any claim 36 to 37 claim 36, characterized in that said vector is an adenoviral vector or a lentiviral vector.
- 39. (original) Pharmaceutical compositions comprising melusin, fragments and/or derivatives thereof for the prevention and/or treatment of heart failure.